
Sequence Listing could not be accepted due to errors.

See attached Validation Report.

If you need help call the Patent Electronic Business Center at (866)

217-9197 (toll free).

Reviewer: Anne Corrigan

Timestamp: [year=2011; month=3; day=22; hr=10; min=46; sec=25; ms=118;]

Reviewer Comments:

<210> 2

<211> 1776

<212> DNA

<213> Artificial Sequence

<220>

<223> modified NS5B

(ERRORED PORTION SHOWN BELOW)

<221> variation

<222> (83) ... (83)

<223> n = G or T

tcnatgtcnt acncntggac nggngccntn atnacaccat gtgggcccga agaggagaag 60 ttaccgatca nccctctgag taattcgctc atncggttcc ataataaggt gtactccaca 120

The above "<220>-<223>" section describing the "n" at location 83 is errored: "a" is at that location. The "n" at location 93 is not explained above.

<210> 4

<211> 4182

<212> DNA

<213> Artificial Sequence

<220>

<223> modified NS3-5A

<221> variation

<222> (2711) ... (2711)

<223> n = T or C

<221> variation

<222> (3645)...(3645)

<223> n = A or G

(ERRORED PORTIONS SHOWN BELOW)

gtgggtccgg gagaggggc tgtgcagtgg atgaaccggc tgatagcgtt cgcctcgcgg 2700 ggtaatcatg tttcccccac gcactatgtg cctgagagcg acgccgcagc gcgtgttact 2760

There is no "n" at location 2711: "t" is at that location.

ttgtctgcgc cttccttgaa ggcgacatgc actacccacc atgtctctcc ggacgctgac 3600 ctcatcgagg ccaacctcct gtggcggcag gagatgggcg ggancatcac ccgcgtggag 3660

There is no "n" at location 3645: "c" is at that location. The "n" at location 3644 is not explained above.

Validated By CRFValidator v 1.0.3

Application No: 10577893 Version No: 1.0

Input Set:

Output Set:

Started: 2011-03-11 10:24:53.976

Finished: 2011-03-11 10:24:56.181

Elapsed: 0 hr(s) 0 min(s) 2 sec(s) 205 ms

Total Warnings: 28

Total Errors: 2

No. of SeqIDs Defined: 28

Actual SeqID Count: 28

Error code		Error Description
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W	213	Artificial or Unknown found in <213> in SEQ ID (2)
E	342	'n' position not defined found at POS: 93 SEQID(2)
W	213	Artificial or Unknown found in <213> in SEQ ID (3)
W	213	Artificial or Unknown found in <213> in SEQ ID (4)
E	342	'n' position not defined found at POS: 3644 SEQID(4)
W	213	Artificial or Unknown found in <213> in SEQ ID (5)
W	213	Artificial or Unknown found in <213> in SEQ ID (6)
W	213	Artificial or Unknown found in <213> in SEQ ID (7)
W	213	Artificial or Unknown found in <213> in SEQ ID (8)
W	213	Artificial or Unknown found in <213> in SEQ ID (9)
W	213	Artificial or Unknown found in <213> in SEQ ID (10)
W	213	Artificial or Unknown found in <213> in SEQ ID (11)
W	213	Artificial or Unknown found in <213> in SEQ ID (12)
W	213	Artificial or Unknown found in <213> in SEQ ID (13)
W	213	Artificial or Unknown found in <213> in SEQ ID (14)
W	213	Artificial or Unknown found in <213> in SEQ ID (15)
W	213	Artificial or Unknown found in <213> in SEQ ID (16)
W	213	Artificial or Unknown found in <213> in SEQ ID (17)
W	213	Artificial or Unknown found in <213> in SEQ ID (18)

Input Set:

Output Set:

Started: 2011-03-11 10:24:53.976

Finished: 2011-03-11 10:24:56.181

Elapsed: 0 hr(s) 0 min(s) 2 sec(s) 205 ms

Total Warnings: 28

Total Errors: 2

No. of SeqIDs Defined: 28

Actual SeqID Count: 28

Error code		Error Description											
W	213	Artificial or Unknown found in <213> in SEQ ID (19)											
W	213	Artificial or Unknown found in <213> in SEQ ID (20) This error has occured more than 20 times, will not be displayed											

SEQUENCE LISTING

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<110> Ludmerer, Steven W.
      Graham, Donald J.
      LaFemina, Robert L.
      Flores, Osvaldo A.
      Pizzuti, Maura
      Traboni, Cinzia
<120> HCV REPLICONS CONTAINING NS5B FROM
  GENOTYPE 2B
<130> 21564YP
<140> 10577893
<141> 2011-03-11
<150> PCT/US2004/036575
<151> 2004-11-03
<150> 60/517,605
<151> 2003-11-05
<160> 28
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<211> 591
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<223> Xaa = threonine or serine
<221> VARIANT
<222> (24)...(24)
<223> Xaa = asparagine or serine
<221> VARIANT
<222> (31)...(31)
<223> Xaa = methionine or isoleucine
<221> VARIANT
<222> (392)...(392)
<223> Xaa = isoleucine or leucine
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1
                                    10
                 5
                                                         15
```

Glu Glu Lys Leu Pro Ile Xaa Pro Leu Ser Asn Ser Leu Xaa Arg

			2.0					0 F					2.0		
Ph≏	Hi⊂	Agn	20 Lvs	Val	Туг	Ser	Thr	25 Thr	Ser	Ara	Ser	ДΊρ	30 Ser	Ţ, _{Q11}	Arα
1110	1110	35	шуы	vai	- y -	DCI	40	TIIT	DCI	1119	DCI	45	DCI	псα	1119
Ala	Lys 50	Lys	Val	Thr	Phe	Asp 55	Arg	Val	Gln	Val	Leu 60	Asp	Ala	His	Tyr
Asp 65	Ser	Val	Leu	Gln	Asp	Val	Lys	Arg	Ala	Ala 75	Ser	Lys	Val	Ser	Ala 80
	Leu	Leu	Thr	Val 85	Glu	Glu	Ala	CÀR	Ala 90	Leu	Thr	Pro	Pro	His 95	
Ala	Lys	Ser	Arg	Tyr	Gly	Phe	Gly	Ala 105	Lys	Glu	Val	Arg	Ser	Leu	Ser
Arg	Arg	Ala 115	Val	Asn	His	Ile	Arg	Ser	Val	Trp	Glu	Asp 125	Leu	Leu	Glu
Asp	Gln 130	His	Thr	Pro	Ile	Asp	Thr	Thr	Ile	Met	Ala 140	Lys	Asn	Glu	Val
Phe 145	Cys	Ile	Asp	Pro	Thr 150	Lys	Gly	Gly	Lys	Lys 155	Pro	Ala	Arg	Leu	Ile 160
Val	Tyr	Pro	Asp	Leu 165	Gly	Val	Arg	Val	Cys 170	Glu	Lys	Met	Ala	Leu 175	Tyr
Asp	Ile	Ala	Gln 180	Lys	Leu	Pro	Lys	Ala 185	Ile	Met	Gly	Pro	Ser 190	Tyr	Gly
Phe	Gln	Tyr 195	Ser	Pro	Ala	Glu	Arg 200	Val	Asp	Phe	Leu	Leu 205	Lys	Ala	Trp
Gly	Ser 210	Lys	Lys	Asp	Pro	Met 215	Gly	Phe	Ser	Tyr	Asp 220	Thr	Arg	Cys	Phe
Asp 225	Ser	Thr	Val	Thr	Glu 230	Arg	Asp	Ile	Arg	Thr 235	Glu	Glu	Ser	Ile	Tyr 240
Gln	Ala	Cys	Ser	Leu 245	Pro	Gln	Glu	Ala	Arg 250	Thr	Val	Ile	His	Ser 255	Leu
		_	260	_		_	_	Pro 265					270	_	
	_	275	_	_		_	280	Ala				285			
	290					295	_	Ile	_		300			_	_
Ala 305	Ala	Gly	Ile	Val	Asp 310	Pro	Val	Met	Leu	Val 315	Cys	Gly	Asp	Asp	Leu 320
Val	Val	Ile	Ser	Glu 325	Ser	Gln	Gly	Asn	Glu 330	Glu	Asp	Glu	Arg	Asn 335	Leu
Arg	Ala	Phe	Thr 340	Glu	Ala	Met	Thr	Arg 345	Tyr	Ser	Ala	Pro	Pro 350	Gly	Asp
Leu	Pro	Arg 355	Pro	Glu	Tyr	Asp	Leu 360	Glu	Leu	Ile	Thr	Ser 365	Cys	Ser	Ser
Asn	Val 370	Ser	Val	Ala	Leu	Asp 375	Ser	Arg	Gly	Arg	Arg 380	Arg	Tyr	Phe	Leu
Thr 385	Arg	Asp	Pro	Thr	Thr 390	Pro	Xaa	Thr	Arg	Ala 395	Ala	Trp	Glu	Thr	Val 400
_				405			_	Leu	410					415	
			420					11e 425					430		
		435		_			440	Gln				445			
_	450		_			455		Leu	_		460				
Arg 465	Leu	His	GLy	Leu	Glu 470	Ala	Phe	Ser	Leu	His 475	Thr	Tyr	Ser	Pro	His 480

Glu Leu Ser Arg Val Ala Ala Thr Leu Arg Lys Leu Gly Ala Pro Pro 485 490 Leu Arg Ala Trp Lys Ser Arg Ala Arg Ala Val Arg Ala Ser Leu Ile 500 505 510 Ala Gln Gly Ala Arg Ala Ala Ile Cys Gly Arg Tyr Leu Phe Asn Trp 515 520 525 Ala Val Lys Thr Lys Leu Lys Leu Thr Pro Leu Pro Glu Ala Ser Arg 530 540 535 Leu Asp Leu Ser Gly Trp Phe Thr Val Gly Ala Gly Gly Gly Asp Ile 550 555 560 545 Tyr His Ser Val Ser His Ala Arg Pro Arg Leu Leu Leu Cys Leu 565 570 575 Leu Leu Ser Val Gly Val Gly Ile Phe Leu Leu Pro Asp Arg 580 585 590

- <210> 2
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- <212> DNA
- <213> Artificial Sequence
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- <223> modified NS5B
- <221> variation
- <222> (3)...(3)
- <223> n = A or T
- <221> variation
- <222> (9)...(9)
- <223> n = C or A
- <221> variation
- <222> (13)...(13)
- <223> n = A or T
- <221> variation
- <222> (15)...(15)
- <223> n = A or C
- <221> variation
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- <223> n A or G
- <221> variation
- <222> (24)...(24)
- <223> n = C or G
- <221> variation
- <222> (28)...(28)
- <223> n = T or C
- <221> modified_base
- <222> (30)...(30)
- <223> n = G or C
- <221> variation

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<221> variation
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<221> variation
<222> (83)...(83)
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<221> variation
<222> (1174)...(1174)
<223> n = A or C
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acctcgagga gtgcctctct gagggcaaag aaggtgactt ttgacagggt gcaggtgctg 180
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tccgtgtggg aggacctcct ggaagaccaa cataccccaa ttgacacaac tatcatggct 420
aaaaatgagg tgttctgcat tgatccaact aaaggtggga aaaagccagc tcgcctcatc 480
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caggettgtt etetgeetea agaageeaga aetgteatae aetegeteae tgagagaett 780
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gagettataa eateetgete eteaaaegta teggtagege tggaeteteg gggtegeege 1140
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tgtggccgct acctcttcaa ctgggcggtg aaaacaaagc tcaaactcac tccattgccc 1620
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<211> 1394
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<222> (1215)...(1215)
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- <221> VARIANT
- <222> (904)...(904)
- <223> Xaa = valine or alanine

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- Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val Glu Gly 20 25 30
- Glu Val Gln Val Ser Thr Ala Thr Gln Ser Phe Leu Ala Thr Cys 35 40 45
- Val Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Ser Lys Thr 50 60
- Leu Ala Gly Pro Lys Gly Pro Ile Thr Gln Met Tyr Thr Asn Val Asp

 70

 75

 80
- Gln Asp Leu Val Gly Trp Gln Ala Pro Pro Gly Ala Arg Ser Leu Thr 85 90 95
- Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg His Ala 100 105 110
- Asp Val Ile Pro Val Arg Arg Gly Asp Ser Arg Gly Ser Leu Leu
 115 120 125
- Ser Pro Arg Pro Val Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro Leu 130 135 140
- Leu Cys Pro Ser Gly His Ala Val Gly Ile Phe Arg Ala Ala Val Cys
 145 150 150
- Thr Arg Gly Val Ala Lys Ala Val Asp Phe Val Pro Val Glu Ser Met 165 170 175
- Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro Pro 180 185
- Ala Val Pro Gln Thr Phe Gln Val Ala His Leu His Ala Pro Thr Gly
 195 200 205
- Ser Gly Lys Ser Thr Lys Val Pro Ala Ala Tyr Ala Ala Gln Gly Tyr 210 220
- Lys Val Leu Val Leu Asn Pro Ser Val Ala Ala Thr Leu Gly Phe Gly 225 230 235
- Ala Tyr Met Ser Lys Ala His Gly Ile Asp Pro Asn Ile Arg Thr Gly
 245 250 255
- Val Arg Thr Ile Thr Thr Gly Ala Pro Val Thr Tyr Ser Thr Tyr Gly 260 270
- Lys Phe Leu Ala Asp Gly Gly Cys Ser Gly Gly Ala Tyr Asp Ile Ile 275 280 285
- Ile Cys Asp Glu Cys His Ser Thr Asp Ser Thr Thr Ile Leu Gly Ile 290 295 300
- Gly Thr Val Leu Asp Gln Ala Glu Thr Ala Gly Ala Arg Leu Val Val 305 310 315 320
- Leu Ala Thr Ala Thr Pro Pro Gly Ser Val Thr Val Pro His Pro Asn 325 330 335
- Ile Glu Glu Val Ala Leu Ser Asn Thr Gly Glu Ile Pro Phe Tyr Gly 340 345 350
- Lys Ala Ile Pro Ile Glu Ala Ile Arg Gly Gly Arg His Leu Ile Phe 355 360 365
- Cys His Ser Lys Lys Cys Asp Glu Leu Ala Ala Lys Leu Ser Gly 370 380
- Leu Gly Ile Asn Ala Val Ala Tyr Tyr Arg Gly Leu Asp Val Ser Val 385 390 395 400

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Thr	Gly	Tyr	Thr		Asp	Phe	Asp			Ile	Asp	Cys			Cys
Val	Thr	Gln	420 Thr	Val	Asp	Phe	Ser	425 Leu	Asp	Pro	Thr	Phe	430 Thr	Ile	Glu
		435					440					445			
Thr	Thr 450	Thr	Val	Pro	Gln	Asp 455	Ala	Val	Ser	Arg	Ser 460	Gln	Arg	Arg	Gly
_	Thr	Gly	Arg	Gly	_	Met	Gly	Ile	Tyr	_	Phe	Val	Thr	Pro	_
465 Glu	Ara	Pro	Ser	Glv	470 Met	Phe	Asp	Ser	Ser	475 Val	T _i eu	Cvs	Glu	Cvs	480 Tvr
	9			485					490			- 1 -		495	- 1 -
Asp	Ala	Gly	Cys 500	Ala	Trp	Tyr	Glu	Leu 505	Thr	Pro	Ala	Glu	Thr 510	Ser	Val
Arg	Leu	Arg	Ala	Tyr	Leu	Asn	Thr		Gly	Leu	Pro	Val		Gln	Asp
		515		_	-: -		520			-: -		525			
His	Leu 530	Glu	Phe	Trp	Glu	Ser 535	Val	Phe	Thr	Gly	Leu 540	Thr	His	Ile	Asp
Ala	His	Phe	Leu	Ser	Gln	Thr	Lys	Gln	Ala	Gly	Asp	Asn	Phe	Pro	Tyr
545 -			_	7	550	_,				555 -				_	560 -
Leu	Val	Ala	Tyr	GIn 565	Ala	Thr	Val	Cys	A1a 570	Arg	Ala	GIn	Ala	9ro 575	Pro
Pro	Ser	Trp	Asp 580	Gln	Met	Trp	Lys	Cys 585	Leu	Ile	Arg	Leu	Lys 590	Pro	Thr
Leu	His	Gly 595	Pro	Thr	Pro	Leu	Leu 600	Tyr	Arg	Leu	Gly	Ala 605	Val	Gln	Asn
Glu	Val 610		Leu	Thr	His	Pro 615		Thr	Lys	Tyr	Ile 620		Ala	Cys	Met
Ser		Asp	Leu	Glu	Val		Thr	Ser	Thr	Trp		Leu	Val	Gly	Gly
625		_			630					635				_	640
Val	Leu	Ala	Ala	Leu 645	Ala	Ala	Tyr	Cys	Leu 650	Thr	Thr	Gly	Ser	Val 655	Val
Ile	Val	Gly	Arg 660	Ile	Ile	Leu	Ser	Gly 665	Arg	Pro	Ala	Ile	Val	Pro	Asp
Arg	Glu	Phe	Leu	Tyr	Gln	Glu	Phe		Glu	Met	Glu	Glu		Ala	Ser
His	Leu	675 Pro	Tyr	Tle	Glu	Gln	680 Glv	Met	Gln	Leu	Ala	685 Glu	Gln	Phe	Lvs
	690		- 1 -			695	1				700				— <i>1</i> ~
Gln 705	Lys	Ala	Leu	Gly	Leu 710	Leu	Gln	Thr	Ala	Thr 715	Lys	Gln	Ala	Glu	Ala 720
	Ala	Pro	Val	Val		Ser	Lys	Trp	Arg		Leu	Glu	Thr	Phe	
				725					730					735	
Ala	Lys	His	Met 740	Trp	Asn	Phe	Ile	Ser 745	Gly	Ile	Gln	Tyr	Leu 750	Ala	Gly
Leu	Ser	Thr 755	Leu	Pro	Gly	Asn	Pro 760	Ala	Ile	Ala	Ser	Leu 765	Met	Ala	Phe
Thr	Ala	Ser	Ile	Thr	Ser	Pro	Leu	Thr	Thr	Gln	Ser	Thr	Leu	Leu	Phe
	770	_	-	-	_	775			-	_	780	_	_	_	
Asn 785	Ile	Leu	Gly	Gly	Trp 790	Val	Ala	Ala	Gln	Leu 795	Ala	Pro	Pro	Ser	Ala 800
Ala	Ser	Ala	Phe		Gly	Ala	Gly	Ile		Gly	Ala	Ala	Val	$\overline{}$	Ser
Ile	Glv	Leu	Glv	805 Lys	Val	Leu	Val	asA	810 Ile	Leu	Ala	Glv	Tvr	815 Gly	Ala
_	∠		820	<u> </u>	_		_	825			- -	<i>_</i>	830	∠	-
Gly	Val	Ala 835	Gly	Ala	Leu	Val	Ala 840	Phe	Lys	Val	Met	Ser 845	Gly	Glu	Met
Pro	Ser		Glu	Asp	Leu	Val		Leu	Leu	Pro	Ala		Leu	Ser	Pro

Gly Ala Leu Val Val Gly Val Val Cys Ala Ala Ile Leu Arg Arg His Val Gly Pro Gly Glu Gly Ala Val Gln Trp Met Asn Arg Leu Ile Ala Phe Ala Ser Arg Gly Asn His Xaa Ser Pro Thr His Tyr Val Pro Glu Ser Asp Ala Ala Arg Val Thr Gln Ile Leu Ser Ser Leu Thr Ile Thr Gln Leu Lys Arg Leu His Gln Trp Ile Asn Glu Asp Cys Ser Thr Pro Cys Ser Gly Ser Trp Leu Arg Asp Val Trp Asp Trp Ile Cys Thr Val Leu Thr Asp Phe Lys Thr Trp Leu Gln Ser Lys Leu Leu Pro Gln Leu Pro Gly Val Pro Phe Phe Ser Cys Gln Arg Gly Tyr Lys Gly Val Trp Arg Gly Asp Gly Ile Met Gln Thr Thr Cys Pro Cys Gly Ala Gln Ile Thr Gly His Val Lys Asn Gly Ser Met Arg Ile Val Gly Pro Lys Thr Cys Ser Asn Thr Trp His Gly Thr Phe Pro Ile Asn Ala Tyr Thr Thr Gly Pro Cys Thr Pro Ser Pro Ala Pro Asn Tyr Ser Arg Ala Leu Trp Arg Val Ala Ala Glu Glu Tyr Val Glu Val Thr Arg Val Gly Asp Phe His Tyr Val Thr Gly Met Thr Thr Asp Asn Val Lys Cys Pro